## 10/581158 (AP20 Rec'd PCT/PTO 31 MAY 2006)

960

## SEQUENCE LISTING <110> Yoshinori Watanabe <120> Novel centromeric protein SHUGOSHIN <130> 4439-4043 <150> JP2003-401943 <151> 2003-12-01 <150> JP2004-279450 <151> 2004-09-27 <160> 45 <170> PatentIn version 3.1 <210> 1 <211> 960 <212> DNA <213> yeast <400> 1 atgaactttc aatttataaa ttcaaatata aacaatgaag ataaattgcc gatggagtcg ttgaaaaaga aatttttaaa acaaaatcgt gaaattataa aaataaatac tcaqctttct 120 ataaaaatta gagaatctga aaacgaaatt caagatttga tacaagaaaa tttcactttg 180 aaaagttatt tggttaaact tgaagctcga tttcgcaatc aatctcaaac tgaggacttg 240 ttaaaaaaact tctttcctga gatacaaacc attcacaaaa agatttcaca agtgcaaagt 300 ttactgaaga ttatagagaa aaagtgttca tcagatttcc tcgaagcgaa tgtaaaaagt 360 caatttacaa cctgtgaaaa taaagattcg aaagaagatt atcagatttt qcataataaa 420 cgcttggagt atgtatcatt taatgatgaa cttaaaagtc tcgaaacagg gcaaccattg 480 tattgttttc aagatttcca aaaaaaagtc catggtcctc cggctctatc tgaaaaacct 540 ggaaaatgta tattaaaaga taaaaccaat gcccacgtaa acaaaatacc acaagatgag 600 gtgaattact cattgccgca aaaaaatatc accatctttt caaaggaatt aaaagaaaac 660 gaatttgaat ccatcaacga gggcgaaact gaagaagaaa aggctaaaac atcaaatgtt 720 tgtgttttgta ttccttgtaa aagtgctgaa cagataactg accttaaagg acaagcaacc 780 ggagacagct ccccatgtga ttttgaagaa tctcaaccaa ggattaatgg acgtgaaaaa 840 ctaagacgat cagtcaaagt gataaactat gcaataccca gtttgcgaac taaactacga 900

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Val Lys Leu Glu Ala Arg Phe Arg Asn Gln Ser Gln Thr Glu Asp Leu 70

Leu Lys Asn Phe Phe Pro Glu Ile Gln Thr Ile His Lys Lys Ile Ser

Gln Val Gln Ser Leu Leu Lys Ile Ile Glu Lys Lys Cys Ser Ser Asp 105

Phe Leu Glu Ala Asn Val Lys Ser Gln Phe Thr Thr Cys Glu Asn Lys 115

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Tyr Cys Phe Gln Asp Phe Gln Lys Lys Val His Gly Pro Pro Ala Leu 165

Ser Glu Lys Pro Gly Lys Cys Ile Leu Lys Asp Lys Thr Asn Ala His 185

Val Asn Lys Ile Pro Gln Asp Glu Val Asn Tyr Ser Leu Pro Gln Lys 195 200

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Asn Ile Asp Leu Arg Thr Thr Ala Ile Asn Leu Glu Glu Gln Leu Glu 50 55 60

Thr Val Gln Asn Glu Asn Glu Glu Asn Lys Thr Lys Leu Ala Ala Leu 70 75 80

Leu Asn Arg Phe His Glu Glu Thr Asp Asn Phe Leu Ser Lys Leu Ser 85 90 95

Leu Cys Gln Gln Glu Ile Gln Asp Thr Phe Lys Pro Val Glu Ala Asn 100 105 110

Leu Ala Tyr Asp Val Asp Thr Asp Ser Glu Asp Leu Asp Glu Glu Ser 115 120 125

Val Val Lys Asp Thr Glu Glu Ile Ile Glu Gln Ala Gln His Asp Val 130 135 140

Ser Leu Arg Asn Leu Ser Gly Ile Glu Asp Glu Asn Ile Ile Asp Asp 145 150 155 160

Gly Glu Thr Ala Ile Asn Glu Gln Lys Lys Arg Glu Ala Asn Val Phe 165 170 175

Ser Asp Thr Gln Ser Ala Pro Gln Leu Lys Ser Gly Lys Ala Leu Pro 180 185 190

Ala Asp Phe Glu Asn Pro Tyr Asn Leu Ser Asn Ser Lys Pro Val Asn 195 200 205

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Ile Asp Ser Ala Pro Gln Glu Lys Asn His Glu Tyr Glu Ile Val Ser225230235240

Pro Lys Ser Leu Ser Asn Lys Ile Asn Asn Gln Ala Ala Gln Arg

÷.

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Met Asp Leu Glu Ser Gln Lys Val Glu Asn Ile Arg Gln Ser Tyr Ser 35 40 45

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Val Asn Ser Leu Glu Lys Lys Ile Ser Gln Leu Val Gln Glu Asn Val 65 70 75 80

Thr Leu Arg Ser Lys Thr Ser Ile Ser Glu Ala Ile Tyr Arg Glu Arg 85 90 95

Leu Ser Asn Gln Leu Gln Val Ile Glu Asn Gly Ile Ile Gln Arg Phe 100 105 110

Asp Glu Ile Phe Tyr Met Phe Glu Asn Val Arg Lys Asn Glu Asn Leu 115 120 125

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Arg Ser Cys Ser Leu Ser Ser Pro Thr Tyr Ser Lys Ser Tyr Thr Arg 145 150 155 160

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Asp Asp Gly Pro Asp Leu Glu Pro Lys Ala Lys Lys Arg Lys Ser Ser 180 185 190

Arg Arg Gln Ser Met Phe Val Ser Thr Ser Leu Glu Pro Glu Asp Glu 195 200 205 Thr Gly Glu Asn Glu Pro Met Met Glu Asn Ser Ser Val Glu Val Pro 210 215 220

Ala Glu Ser His Glu Ser Ala Gln Val Glu Glu Thr Ile Asp Ala Leu 225 230 235 240

Asn Pro Glu Glu Asn Ser Asp Ser Val Ser Asn Phe Thr Asn Ser 245 250 255

Ile Ile Glu Tyr Ser Ile Pro Glu Glu Asn Pro Thr Glu Pro Glu His  $260 \\ \hspace{1.5cm} 265 \\ \hspace{1.5cm} 270 \\ \hspace{1.5cm}$ 

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Thr Val Pro Ser Asn Pro Leu Pro Leu Pro Leu Pro Gly Pro Ser Ala 290 295 300

Thr Leu Pro Thr Thr Ser Asp Ala Ser Thr Val Tyr Pro Ser Ser 305 310 315 320

Ser Ser Ser Thr Asn Ser His Pro Lys Thr Lys Ile Lys His Ser Met 325 330 335

Lys Pro Pro Arg Ile Glu Leu Lys Lys Lys Val Ile Asp Glu Val Met 340 345 350

Pro Val Ser Asn Met Ser Ser Asn Ser Glu Ile Ser Phe Thr Arg Thr 355 360 365

Arg Arg Thr Arg Gly Lys Ala Val Asp Tyr Thr Leu Pro Ser Leu Arg 370 375 380

Ala Lys Met Arg Arg Pro Ser Glu Lys Leu Val Asp Ala Thr Thr Val 385 390 395 400

Ile Asp Ile His Asp Leu Gln Val Ser Lys Arg Asn Arg Glu Thr Ser  $405 \hspace{1.5cm} 410 \hspace{1.5cm} 415$ 

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Met	Ala	Arg	Leu	Asn	Glu	Gln	Ala	Met	Ser	Ser	Val	Ala	Leu	Ser	Thr
1				5					10					15	

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Ile Ala Arg Val Asn Ser Thr Gln Ser Leu Arg Ile Arg Gly Leu Glu 35 40 45

Asn Glu Cys Ala Arg Leu Leu Ser Glu Asn Leu Glu Leu Arg Gly Gln 50 55 60

Val Leu Arg Leu Glu Lys Glu Leu Gln Asp Asn Ala Ala Arg Arg Val 65 70 75 80

Ala Asp His Ala Leu Glu Val Lys Ala Lys Met Glu Thr Gln Leu Ala 85 90 95

Glu Leu Ser Ser Leu Leu Ala Ser Leu Gly Glu Pro Pro Ser Lys Arg 100 105 110

Arg Leu Ser Glu Glu Arg Arg Tyr Ala Gln Pro Arg Pro Ser Val His 115 120 125

Arg Ser Pro Pro Leu Arg Arg Ala Arg Gln Glu Ala Asp Gln Glu Leu 130 135 140

Leu Ala Glu Gln Glu Gly Arg Leu Pro Pro İle Tyr Glu Asn Lys Thr 145 150 155 160 Tyr Ala Arg Ala Thr Met Asn Ser Glu Glu Ile Leu Ala Leu Cys Met 165 170 175

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Ser Arg Phe Val Glu Asp Asp Met Val Ile Pro Cys Ser Pro Ser Pro 195 200 205

Asn Lys Asn Ala Glu Ala Glu Glu Thr Glu Thr Thr Glu Gln Val Glu 210 215 220

Glu Ser Pro Arg Ala Leu Gln Val Pro Pro Ser Leu Ser Pro Pro Lys 225 230 235 240

Leu Asp Tyr Asp Arg Arg Pro Asn Met Ile Leu Phe Ser Pro Pro Lys 245 250 255

Glu Ser Arg Val Ala Glu Pro Ser Lys Met Phe Ser Pro Pro Pro Met 260 265 270

Glu Pro Pro Lys Gln Ser Thr Ser Ala Val Pro Ser Glu Thr Ile Arg 275 280 285

Ala Gly Leu Lys Arg Lys Leu Asn Gly Asp Asn Gln Asn Glu Pro Asn 290 295 300

Lys Ala Thr Lys Leu Gln Gln Gly Lys Glu Asn Gly Asn Glu Thr Gly 305 310 315 320

Ile Lys Lys Gly Leu Ser Ala Arg Asp Pro His Lys Arg Lys Ser Ile 325 330 335

Lys Glu Thr Ala Thr Lys Pro Arg Ala Pro Leu Ser Ala Lys Ser Thr 340 345 350

Asn Glu His Ile Val Ser Pro Lys Lys Pro Ala Lys Pro His Gln Val 355 360 365

Ala Asp Asp Phe Lys Pro Val Lys Val His Lys Ala Ser Lys Gly Lys 370 380

Glu Lys Val Asp Leu Pro Ala Pro Asp Lys Lys Ser Ala Val Glu Glu 385 390 395 . 400

Thr Gln Gly Asn Ser Thr Ser Ala Phe Thr Lys Val Glu Ile Leu Pro  $405 \hspace{1.5cm} 410 \hspace{1.5cm} 415 \hspace{1.5cm}$ 

Pro Ala Leu Glu Pro Thr Pro Glu Val Ala Glu Ile Pro Glu Thr Asp 420 425 430

Ile Leu Ile Thr Pro Gly Thr Pro Glu Arg Ala Ser Glu Ser Thr Val435  $\phantom{0}440$   $\phantom{0}445$ 

Val Thr His Asp Thr Pro Pro Pro Ala His Ile Ser Ser Asn Gly Glu 450 455 460

Thr Ser Arg Pro Ser Arg Arg Ala Arg Ala Ala Ile Ser Tyr Thr Glu 465 470 475 480

Pro Asn Leu Arg Asp Lys Met Arg Arg Pro Thr Lys Glu Leu Phe Asp 485 490 495

Ala Val Ser Gly Glu Gly Lys Phe Leu His Arg Pro Thr Ser Gln Gln 500 505 510

Gln Gln Gln Arg Lys Gly Asp Glu Ser Ala Pro Thr Ser Val Ser 515 520 525

Lys Val Lys Val Glu Pro Ser Pro Ala Val Asp Ile Ser Ser Leu Thr 530 540

Ser Ser Ala Leu Phe Glu Lys Glu Lys Glu Lys Glu Pro Gln Pro Asp 545 550 555 560

Glu Gly Ile Leu Ser Pro Asn Gly Ile Leu Pro Ser Ser Val Asp Leu 565 570 575

Gly Arg Arg Arg Ala Ser Ser Phe Ser Thr Ala Ala Pro Ala Met 580 585 590

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Leu Ser Asn Ser Ile Thr Thr Arg Pro Arg Gly Gly Lys Gly Arg Gln 630 635

Ser Met Ser Arg Ser Val Pro Thr Ile Pro Thr Glu Asn Tyr Glu His 645

Glu Asp Ala Gln Leu Ser Thr Asn Ser Ala Ser Val Asp Leu Tyr Asp 665

Phe Ala Ser Cys Ala Ser Pro Asp Ser Ala Ala Pro Gln Leu Glu Ala 675 680

Thr Thr Gly Asp Val Pro Val Asn Lys Lys Ala Pro Lys Gly Ser Arg 695 700

Arg Ala Ser Ser Ala Ala Ser Thr Glu Thr Thr Ala Thr Ala Ser Ala 710 720

Lys Pro Arg Ser Ser Arg Lys Arg Ala Ser Met Leu Val Pro Lys Lys 725 730 735

Ser Leu Trp Ala Glu Glu Leu Ala Gln Glu Glu Asp Glu Glu Asp 740 745

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Met Asn Ser Ala Gln Arg Arg Lys Leu Gly Asp Ile Thr Asn Leu Gln 35 40 45

Asn Gln Lys Asn Leu Met Asn Gln Gly Ala Lys His Gln Gln Gln Ala 50 60

Ile Leu Ile Ser Ser Lys Glu Asn Ala Glu Asn Leu Gln Lys Ala Leu 65 70 75 80

Arg Asn Ser Ser Glu Asn Thr Lys Leu Met Lys Val Val Met Glu Arg 85 90 95

Asp Gly Ile Lys Ser Asp Leu Lys Lys Leu Arg Ile Glu Phe Gln Lys
100 105 110

Val Gln Glu Gln Asn Leu Leu Leu Ala Gln Ala Asn Thr Arg Ile Leu 115 120 125

Ala Leu Lys Val Leu Gln His Glu Leu Gly Cys Lys Asn Gly Leu Val 130 135 140

Met Ala Arg Lys Met Leu Leu Lys Ala Gln Ala Asn Ala Cys Gly Gly 145 150 155 160

Ala Cys Lys Thr Phe Gln Pro Asn Asp Ala Asp His Glu His Ala Ser 165 170 175

Gly Ser Ser Asn Ala Asn Ser Leu Gln Arg Asn Glu Lys Ala Asn Ser 180 185 190

Lys Arg Arg Val Ser Gly Arg Lys Asn Pro Ala Asn Ser Glu Val Leu 195 200 205

Asp Ile Ile Gly Arg Ser Gly Glu Thr Cys Gln Met Glu Asp Asn Ile 210 215 220

Asp Asn Lys Lys Leu Val Ser Asp Ser Asp Asn Asp Ala Glu Asn His 225 230 235 240

Ile Asn Asp Asn Val Gln Ser Lys Arg Tyr Cys Ala Gly Arg Gln Ser 245 250 255

Ser Ser Ser Lys Thr Arg Glu Ala Ser Gln Thr Glu Thr Leu Gln Lys 260 265 270

Val Val Asp Ala Lys Glu Ile Lys Gly Asp Ala Arg Phe Ser Leu Thr 275 280 285

Lys His Ser Asp Trp Leu Lys Ser Gln Glu Pro Glu Pro Ser Glu Ser 290 295 300

Leu Tyr Glu Ser Arg Phe Pro Leu Arg Arg Ser Ala Arg Leu Lys 305 310 315 320

Ser Gln Glu Pro Glu Pro Ser Glu Ser Phe His Asp Ser Ile Glu Thr .  $325 \hspace{1.5cm} 330 \hspace{1.5cm} 335$ 

Thr Lys Arg Arg Arg Ser Ala Ile Arg Ser Ala Met Phe Asn Ile Gln 340 345 350

Glu Leu Gly Val Ile Gln Asn Leu Asn Gly Leu Pro Asp Asp Gln Glu 355  $\phantom{\bigg|}360\phantom{\bigg|}365\phantom{\bigg|}$ 

Ile Ala Ala Lys Ala Arg Cys Ser Ala Arg Glu Gln Ser Thr Gly Ser 370 375 380

Lys Pro Glu Ala Val Glu Pro His Asp Thr Lys Glu Ile Ile Gly Lys 385 390 395 400

Ser Arg Ile Ser Leu Arg Arg Gln Ser Ala Arg Phe Asn Phe Gln Glu 405 410 415

Leu	Gly	Val	Thr 420	Glu	Asn	Leu	Asn	Gly 425	Pro	His	Asp	Asp	Gln 430	Thr	Ile	
Ala	Ala	Asn 435	Ala	Arg	Cys	Cys	Ala 440	Ser	Glu	Gln	Ser	Ile 445	Gly	Ser	Lys	
Pro	Glu 450	Ala	Val	Glu	Pro	His 455	Asp	Ile	Glu		Arg 460	Ile	Gly	Lys	Ile	
Arg 465	Val	Ser	Ser	Arg	Arg 470	Gln	Ser	Ala	Asn	Ile 475	Glu	Thr	Pro	Arg	Ala 480	
Ile	Lys	Glu	Pro	Ala 485	Asn	Pro	Pro	Leu	His 490	Asp	Asp	Asn	Val	Glu 495	Glu	
Ser	Ser	Gln	Ile 500	Ser	Cys	Ser	Val	Ser 505	Met	Glu	Leu	Lys	Arg 510	Glu	Ser	
Lys	Lys	Lys 515		Thr	Gly	Asp	Glu 520	Ser	Glu	Glu	Met	Arg 525	Lys	Thr	Thr	
Val	Gly 530	Arg	Pro	Ser	Arg	Gln 535	Ala	Ala	Glu	Lys	Ile 540	Lys	Ser	Tyr	Lys	
Glu 545	Pro	Ser	Leu	Lys	Glu 550	Lys	Met	Arg	Gly	Gly 555	Phe					
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gcto	ctago	cac a	accga	aaata	aa a	ctcgt	cga	g tto	gagco	ggta	ttga	agatt	ca (	gaaac	tgagg	180
atta	aactt	tac q	ggagt	gtgo	ca go	gaaaa	agaat	t ttq	gcago	cttg	ctca	aggca	aaa o	cagto	agatg	240
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aaac	catct	ttg a	agga	gcaaq	gt ad	cttco	cacgt	t aca	acato	catg	aato	cgaaa	aga (	caago	ıtttca	360

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Gln Glu Tyr Ala Ala Lys Leu Gln Lys Ala Phe Pro Leu His Phe Asn 20 25 30

Leu Glu Asn Met Thr Leu Met Lys Ala Leu Ala His Arg Asn Lys Leu  $35 \hspace{1cm} 40 \hspace{1cm} 45$ 

Val Glu Leu Ser Gly Ile Glu Ile Gln Lys Leu Arg Ile Asn Leu Arg 50 55 60

Ser Val Gln Glu Lys Asn Leu Gln Leu Ala Gln Ala Asn Ser Gln Met 65 70 75 80

Leu Ala Leu Lys Asp Leu Gl<br/>n His Glu Leu Gly Cys Lys As<br/>n Ala Leu 85 90 95

Leu Lys Val Lys Lys His Leu Glu Glu Gln Val Leu Pro Arg Thr His  $100 \\ 105 \\ 110$ 

His Glu Ser Lys Asp Lys Val Ser Ala Ser Ala Ser Asp Gly Asp Cys 115 120 125

Lys Ser Phe Gln Val His Asp Ile Lys His Lys Asp Thr Lys Arg Lys 130 135 140

Arg Thr Thr Arg Ile Lys Ser Ser Val Ser Ala Asp Val Lys Pro Ile 145 150 155 160

Pro Val Asn Asp Ser Asn Ser Lys Ala Asn Arg Lys Arg Arg Val Ser 165 170 175

Gly Val Ile Asp Thr Thr Gly Ile Pro Glu Glu Ile Cys Gln Thr Glu
180 185 190

Asp Asp Ile Asp Lys Gly Val Val Ser Arg Gly Val Asn Gln Asp Ile 195 200 205

Asp Asn Val Val Asn Lys Lys Phe Val Pro Asp Ala Ala Asn Pro Val 210 215 220

Lys Glu Ser Val His Arg Lys Arg Gln Cys Thr Arg Arg Gln Ser Thr 225 230 235 240

Arg Phe Asp Val Gln Glu Thr Lys Gln Thr Glu Lys Leu Leu Glu Met 245 250 255

Asp Gly Ala Lys Glu Ser Lys Glu Thr Ala Ser Phe Ser Leu Arg Arg 260 265 270

Arg Ser Ala Arg Leu Arg His Glu Glu Ala Glu Pro Cys Lys Ser Leu 275 280 285

His	Glu 290	Gly	Asp	Glu	Val	Arg 295	Glu	Thr	Ile	Lys	Arg 300	Arg	Arg	Val	Ser		
Leu 305	Arg	Leu	Ser	Ala	Arg 310	Phe	Asp	Ile	Gln	Glu 315	Pro	His	Val	Thr	Glu 320		
Thr	Ser	Asn	Ala	Asp 325	Asp	Ala	Arg	Ser	Ile 330	Val	Ile	Glu	Glu	Ser 335	Ala		
Gly	Ser	Arg	Ser 340	Glu	Ser	Val	Glu	Pro 345	Ser	Glu	Ser	Arg	His 350	Glu	Thr		
Lys	Glu	Ile 355	Thr	Arg	Lys	Arg	Ser 360	Phe	Ser	Thr	Arg	Arg 365	Gln	Ser	Thr		
Lys	Gly 370	Lys	Ser	Gln	Thr	Asp 375	Glu	Ala	Ile	Lys	Glu 380	Ile	Ala	Thr	Asp		
Pro 385	Ser	Leu	Val	Asn	Thr 390	Ile	Val	Gln	Glu	Cys 395	Asp	Gln	Glu	Thr	Glu 400		
Ser	Lys	Asp	Lys	Pro 405	Lys	Ala	Asp	Glu	Asn 410	Glu	Gly	Met	Thr	Arg 415	Arg		
Ser	Ser	Val	Gly 420	Arg	Pro	Ser	Arg	His 425	Ala	Ala	Glu	Lys	Val 430	Gln	Ser		
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gtca	atcc	tgc a	aact	gaga	aa a	gaat	gcta	c tad	cctt	actt	gtca	agcto	gta	tgcat	tgaaa	3	300

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<211> 517

<212> PRT

<213> mosue

<400> 14

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Ile Gly Lys Arg Lys Ser Phe Ile Val Ala Pro Gly Gln Val Pro Thr 35 40 45

Asn Thr Ala Thr Leu Leu Arg Tyr Tyr Gln Asp Asn Asn Arg Leu Leu 50 55 60

Val Leu Ala Leu Glu Asn Glu Lys Ser Lys Val Arg Glu Ala Gln Asp 65 70 75 80

Val Ile Leu Gln Leu Arg Lys Glu Cys Tyr Tyr Leu Thr Cys Gln Leu  $85 \hspace{1cm} 90 \hspace{1cm} 95$ 

Tyr Ala Leu Lys Glu Lys Leu Thr Ser Arg Gln Ser Glu Glu Thr Thr 100 105 110

Gln Asn Trp Lys Gly Arg Pro Ser Asp Val Val Ser Ser Ile Asp Asn 115 120 125

Thr Thr Arg Asp Leu Ser Gly Lys Ser Leu Gln Gln Ile Ala Val Glu 130 135 140

Glu Thr Asp Cys Pro Tyr Gln Thr Thr Glu Pro Ser Pro Ala Val Thr 145 150 155 160

Pro Glu Thr Gln Gly Cys Asp Phe Asp Ser Gly Lys Val Glu Ser Thr \$165\$ \$170\$ \$175\$

Asp Glu Val Leu Pro Arg Thr Ile Ser Ile Arg Arg His Leu Arg Lys 180 185 190

Asp Phe Ser Asn Ile Ser His Ser Thr Thr Leu Glu Asp Cys Lys Ala 195 200 205

Ser Pro Arg Val Ala Gln Ser Leu Glu Val Lys Gly Ser Arg Cys Arg 210  $\,$  215  $\,$  220  $\,$ 

Glu Val Thr Val Thr Leu His Arg Leu Glu Asn Val Cys Leu Trp Asn 225 230 235 240

Lys Asp Gln Ile Ser Leu Cys Ser Arg Leu Ile Asn Pro Ala Lys Ile 245 250 255

Thr Glu Thr Glu Val Ile Leu Ser Ser Lys Pro Glu Gln Ile Glu Ser 260 265 270

Lys His Lys Arg Ala Arg Lys Arg Arg Ala Glu Gln Arg Arg Thr Lys 275 280 285

Gln Arg Cys Lys Ser Lys Ser Ser Leu Arg Ser Lys Gly Asn Lys Asn 290 295 300

Lys Asp Lys Gln Gly Leu Pro Pro Thr Thr Leu Asp Gly Gly Ile Gly 305 310 315 320

Ser Cys Asp Ala Tyr Asp Phe Asn Leu Lys Gly Thr Val His Pro Thr 325 330 335

Pro Phe Arg Gln Lys Met Asn Asn Gly Cys Asn Lys Glu Thr Asp Ser 340 345 350

Ser Asn Ser Glu Val Ser Asp Leu Glu Cys Ser Thr Ser Glu Asp Glu 355 360 365

Ser Asp Asp Leu Tyr Leu Pro Pro Ser Lys Arg Leu Arg Asp Tyr Arg 370 375 380

Glu Ser Glu Arg Ala Val Thr Arg Pro Arg Ser Lys Arg Gly Leu Gln 385 390 395 400

Thr Gly Ile Pro Pro Glu Thr Gln Glu Ser Pro Arg Cys Ser Leu Lys 420 425 430

Asp Val Thr Asn Ile Leu Gln Cys Pro Arg Val Lys Ile Arg Lys Pro 435 440 445

Ser Leu Pro Pro Lys Arg Arg Glu Asp Ser Pro Ala Val Ala Leu Thr 450 455 460

Lys Arg Arg Cys Ser Thr Ile Lys Ser Tyr Lys Glu Pro Thr Leu Ala 465 470 475 480 Ser Lys Leu Arg Arg Gly Asp Pro Phe Thr Asp Leu Cys Phe Leu Asn 485 490 495

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Thr Lys Gln Thr Gln 515

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<211> 3495

<212> DNA

<213> mouse

<400> 15

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Met Glu Tyr Pro Gly Ile Lys Val Asp Thr Val Thr Ser Gly Ile Gln

5 \_\_10 \_\_15

Arg Arg Val Lys Gly Arg Ile Ala Lys Thr Asn Leu Asn Val Ser Leu 20 25 30

Ala Ser Lys Ile Lys Ala Lys Ile Leu Asn Asn Ser Ser Ile Phe Lys 35 40 45

Ile Ser Leu Lys His Asn Asn Arg Ala Leu Ala Arg Ala Leu Ser Lys 50 55 60

Glu Lys Glu Asn Ser Arg Arg Ile Thr Thr Glu Lys Met Gln Leu Gln 65 70 75 80

Lys Glu Val Glu Lys Leu Asn Phe Glu Asn Thr Phe Leu Arg Leu Lys 85 90 95

Leu Asn Thr Leu Asn Lys Lys Leu Val Glu Ile Glu Ser His Val Ser 100 105 110

- Asn Asp Leu Leu Thr Ala Ile Glu Ile Ser Ser Leu Ser Glu Phe His 115 120 125
- Gln Gly Ser Phe Leu Leu Ser Ala Thr Lys Lys Gln Arg Asn Ser Lys 130 135 140
- Gln Cys Lys Pro Ala His Leu Pro Tyr Ala Arg Val Leu Leu Thr Ser 145 150 155 160
- Glu Asn Asp Asp Asp Gly Ala Asp Asp Lys Trp Gln Thr Lys Cys 165 170 175
- Asn Asn Arg Thr Ile Ser Lys Thr Ser Pro Asp Ser Thr Ser Ser Val 180 185 190
- Ser Arg Gln Pro Ser Ser Leu His Gln Cys Asn Leu Lys Ala Phe Pro 195 200 205
- Pro Lys Glu Asp Asn Gln Lys Thr Cys Gly Ser Gly His Leu Glu His 210 215 220
- Thr Ser Ser Val Asp Ile Leu Pro Asn Glu Ser His Ser Asp Gln Ser 225 230 235 240
- Pro Lys Ser Ser Leu Ser Glu Met Lys Thr Ala Pro Ser Pro Ser Leu 245 250 255
- Arg Arg Glu Lys Leu Ser His Gly Asn Val Thr Met Arg Lys Lys Cys 260 265 270
- Val Ser Ser Thr Pro Asp Ile Leu Tyr Val Thr Asp Leu Asp His Gln 275 280 285
- Pro Thr Ser Ser Pro Gly Ser Asn Trp Asn Asn Glu Ile His Gly His 290 295 300
- Thr Asn Glu Thr Ser Asn Asn Thr Gln Arg Asn Ala Glu Cys Phe Leu 305 310 315 320
- Asp Leu Pro Ser Glu Ser Ser Ser Glu Pro Asp Ala Lys Arg Met Glu 325 330 335

Leu Val Gln Lys Asn Thr Asp Ser Phe His Phe Gln Lys Thr Val Tyr 340 345 350

Asp Ala Ala Asp Met Glu Leu Thr Ala Thr Asp Ile Gly Lys Ile Val 355 360 365

Ala Val Ser Lys Ser Lys Lys Asn Gln Asn Lys Lys Lys Ala Asp Cys 370 375 380

Arg Lys Glu Thr Phe Arg Lys Val Lys Gly Ala Ser Ser Asp Lys Lys 385 390 395 400

Arg Glu Ser Ser Lys Arg Glu Cys Lys Asp Gly Ser Glu Val Gly Ala 405 410 415

Glu Glu Glu Ala Asp Ala Ala Arg Ala Glu Arg Gly Ala Gly Val Leu 420 425 430

Asp Gly Arg Gly Asp Ser Glu Glu Pro Asn Cys Ile Ser Ser Thr Glu 435 440 445

Gln Pro Ser Gln Val Asn Thr Gln Lys Lys Arg Thr Leu Gln Asn Ser 450 460

Ser Asp Gln Glu Asn Ile Gln Asn Thr Lys Arg Arg Gln Thr Tyr Thr 465 470 475 480

Thr Asp Glu Glu Glu Thr Asn Pro Phe Ser Arg His Ser Val Lys \$485\$

Phe Leu Gln Asp Gly Lys Phe Asp Leu Cys Gln Lys Thr Leu His His 500 505 510

Asn Leu Ser Lys Pro Ser Arg Gln Thr Phe Val Ile Arg Lys Ser Glu 515 520 525

Lys Asp Asn Leu Phe Pro Asn Gln Glu Asp Lys Asp Thr Ile Ser Glu 530 535 540

Asn Leu Glu Val Thr Asn Glu Phe His Ile Asp Asp Leu Ser Ile Glu 545 550 555 560

Ala Asn Glu Asn Val Cys Asp His Glu Thr Gln Thr Met Leu Asp Leu

565 570 . 575

Lys Lys Ser Val Ser Ala Gln Gln Asn Gln Thr Lys Ile Asn Lys Thr 580 595

Lys Gln Lys Ile Asn Arg Arg Thr Lys Ile Ile Ser Val Met Ser Gln 595 600 605

Val Tyr Glu Asp Asn Asp Lys Asp Ile His Val Leu Glu Lys Asp Asn 610 615 620

Phe Pro Phe His Thr Gln Ala Asn Lys Glu Thr Thr Ser Gly Asn Leu 625 630 635 640

Glu Ser Ser Lys Glu Phe Glu Ser Pro Leu Leu Phe Thr Arg Asp Asn
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650
655

Gly Ser Leu Arg Asp Cys Lys Thr Gln Asn Val Leu Asp Leu His Lys 660 665 670

Gln Ile Pro Asp Leu Tyr Pro Asp Arg Asn Glu Ser Gln Ile Ser Lys 675 680 685

Lys Cys Phe Ser Asn Asp Gln Gly Val His Cys Ser Glu Lys Asp Lys 705 710 715 720

Ser Leu Leu Gln Lys Asp Lys Asp Phe Pro Gly Thr Leu Lys Asp 725 730 735

Leu Ser Glu Phe Asp Thr Pro Ala Phe Cys Asn Lys Asp Ser Ala Lys 740 745 750

Ser Cys Asp Tyr Lys Ser Glu Met Leu Leu Gly Leu Lys Lys His Asp 755 760 765

Pro Asn Met Gln Pro Ala Cys Gln Asp Asp Ser Lys Ala Gly Lys Lys 770 775 780

Leu Arg Gln Lys Val Asn Arg Lys Thr Glu Ile Ile Ser Lys Ile Thr 785 790 795 800

- Gln Ile His Glu Asn Asp Arg Gly Ser Thr His Asp Ser Leu Asn Lys 805 810 815
- Lys Leu Cys Gln Lys Val Asn Ile Ser Lys Ile Ile Ser Gln Met Asn 820 825 830
- Gln Ile Tyr Glu Thr Ile Asn Glu Asp Gly Asn Gly Phe Lys Ser Ser 835 840 845
- Ile Lys Asp Cys Glu Asp Ile Lys Ser Cys Asp Phe Gly Glu Ile Asn 850 855 860
- Ser Asn Lys Lys Glu Asn Tyr Asp Pro Ile Gln Asp Pro Cys Thr Leu 865 870 875 880
- Val Lys Lys Thr Lys Arg Lys Gly Ser Cys Lys Ala Gly Ser Ser Leu 885 890 895
- Ala Gly Ala Lys Asn Arg Cys Gly Leu Gln Leu Thr Asp Ser Ser Gln
  900 905 910
- Val Gln Ser Val Pro Leu Asp Ser Gly Leu Arg His His Pro Asn Glu 915 920 925
- Ala Asp Ser Gly Pro Gly Glu Gln Thr Asn Leu Pro Lys Met Gln Lys 930 935 940
- Gln Ser Ala Gly Arg Ser Leu Gly Asp Ala Phe Ser Val Ser Leu Gly 945 950 955 960
- Lys Glu Gly Ser Arg Pro Ala Lys Ala Val Ser Lys Met Thr Pro Lys 965 970 975
- Ser Lys Lys Arg Lys Leu Pro Leu Gly Cys Ser Pro Glu Thr His Gly 980 985 990
- Thr Val Glu Ile Thr Pro Asn Thr Asp Leu Ala Lys Ala Val Asp Ser 995 1000 1005
- Gln Gln Thr Glu Lys Glu Asn Tyr Leu Glu Lys Glu Lys Ile Ala 1010 \$1015\$

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Glu	Thr 1040	Cys	Ser	Ser	Asn	Ile 1045	Lys	Asn	Ser	Ser	Leu 1050	Asp	Ser	Met	
Cys	Lys 1055	Ser	Ser	Leu	Pro	Leu 1060	Ser	Ile	Ser	Ser	Arg 1065	Lys	Thr	Leu	
Met	Leu 1070	Glu	Glu	Ser	Ser	Ser 1075	Leu	Glu	Ser	Thr	Cys 1080	Ile	Phe	Gln	
Val	Gly 1085	Asp	Ala	Ala	His	Glu 1090		Ile	Thr	Thr	Gly 1095	Thr	Arg	Asn	
Pro	His 1100	His	Arg	Thr	Gln	Lys 1105		Thr	Pro	Gly	Ser 1110	Arg	Thr	Ser	
Leu	Val 1115	Leu	Val	Asp	Thr	Ser 1120		Val	Ser	Asp	Thr 1125	Asn	Pro	Ala	
Asn	Pro 1130	Glu	Asn	Glu	Ser	Glu 1135	Gly	Gln	Ser	Ser	His 1140	Pro	Met	Arg	
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Arg	Ser 1160	Lys	Met	Arg	Arg										
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											_			ccaaga	180
														cattgaa	240
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<211> 511

<212> PRT

<213> Homo sapiens

<400> 18

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- Val Leu Ala Leu Glu Asn Glu Lys Ser Lys Val Lys Glu Ala Gln Asp 50 55 60
- Ile Ile Leu Gln Leu Arg Lys Glu Cys Tyr Tyr Leu Thr Cys Gln Leu 65 70 75 80
- Tyr Ala Leu Lys Gly Lys Leu Thr Ser Gln Gln Thr Val Glu Pro Ala 85 90 95
- Gln Asn Gln Glu Ile Cys Ser Ser Gly Met Asp Pro Asn Ser Asp Asp 100 105 110
- Ser Ser Arg Asn Leu Phe Val Lys Asp Leu Pro Gln Ile Pro Leu Glu 115 120 125
- Glu Thr Glu Leu Pro Gly Gln Gly Glu Ser Phe Gln Ile Glu Asp Gln 130  $$135\$
- Ile Pro Thr Ile Pro Gln Asp Thr Leu Gly Val Asp Phe Asp Ser Gly
  145 150 155 160
- Glu Ala Lys Ser Thr Asp Asn Val Leu Pro Arg Thr Val Ser Val Arg 165 170 175
- Ser Ser Leu Lys Lys His Cys Asn Ser Ile Cys Gln Phe Asp Ser Leu 180 185 190
- Asp Asp Phe Glu Thr Ser His Leu Ala Gly Lys Ser Phe Glu Phe Glu 195 200 , 205
- Arg Val Gly Phe Leu Asp Pro Leu Val Asn Met His Ile Pro Glu Asn 210 215 220
- Val Gln His Asn Ala Cys Gln Trp Ser Lys Asp Gln Val Asn Leu Ser 225 230 235 240
- Pro Lys Leu Ile Gln Pro Gly Thr Phe Thr Lys Thr Lys Glu Asp Ile 245 250 255

Leu Glu Ser Lys Ser Glu Gln Thr Lys Ser Lys Gln Arg Asp Thr Gln 260 265 270

Glu Arg Lys Arg Glu Glu Lys Arg Lys Ala Asn Arg Arg Lys Ser Lys 275 280 285

Arg Met Ser Lys Tyr Lys Glu Asn Lys Ser Glu Asn Lys Lys Thr Val 290 295 300

Pro Gln Lys Lys Met His Lys Ser Val Ser Ser Asn Asp Ala Tyr Asn 305 310 315

Phe Asn Leu Glu Glu Gly Val His Leu Thr Pro Phe Arg Gln Lys Val 325 330 335

Ser Asn Asp Ser Asn Arg Glu Glu Asn Asn Glu Ser Glu Val Ser Leu 340 345 350

Cys Glu Ser Ser Gly Ser Gly Asp Asp Ser Asp Asp Leu Tyr Leu Pro 355 360 365

Thr Cys Lys Tyr Ile Gln Asn Pro Thr Ser Asn Ser Asp Arg Pro Val 370 375 380

Thr Arg Pro Leu Ala Lys Arg Ala Leu Lys Tyr Thr Asp Glu Lys Glu 385 390 395 400

Thr Glu Gly Ser Lys Pro Thr Lys Thr Pro Thr Thr Thr Pro Pro Glu 405 410 415

Thr Gln Gln Ser Pro His Leu Ser Leu Lys Asp Ile Thr Asn Val Ser 420 425 430

Leu Tyr Pro Val Val Lys Ile Arg Arg Leu Ser Leu Ser Pro Lys Lys 435 440 445

Asn Lys Ala Ser Pro Ala Val Ala Leu Pro Lys Arg Arg Cys Thr Ala 450 455 460

Ser Val Asn Tyr Lys Glu Pro Thr Leu Ala Ser Lys Leu Arg Arg Gly 470 475 480

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1320

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<213> Homo sapiens

<400> 20

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Val Ser Leu Ala Ser Lys Ile Lys Thr Lys Ile Leu Asn Asn Ser Ser 40

Ile Phe Lys Ile Ser Leu Lys His Asn Asn Arg Ala Leu Ala Gln Ala 55

Leu Ser Arg Glu Lys Glu Asn Ser Arg Arg Ile Thr Thr Glu Lys Met 70 75

Leu Leu Gln Lys Glu Val Glu Lys Leu Asn Phe Glu Asn Thr Phe Leu 85 90

Arg Leu Lys Leu Asn Asn Leu Asn Lys Lys Leu Ile Asp Ile Glu Ala 100 105 110

Leu Met Asn Asn Asn Leu Ile Thr Ala Ile Glu Met Ser Ser Leu Ser 115 120 125

Glu Phe His Gln Ser Ser Phe Leu Leu Ser Ala Ser Lys Lys Arg 130 135 140

Ile Ser Lys Gln Cys Lys Leu Met Arg Leu Pro Phe Ala Arg Val Pro 145 150 155 160

Leu Thr Ser Asn Asp Asp Glu Asp Glu Asp Lys Glu Lys Met Gln Cys 165 170 175

Asp Asn Asn Ile Lys Ser Lys Thr Leu Pro Asp Ile Pro Ser Ser Gly 180 185 190

Arg Thr Thr Gln Pro Leu Ser Thr Gln Asp Asn Ser Gly Val Leu Phe 195 200 205

Leu Lys Glu Asn Asn Gln His Val Tyr Gly Leu Asp Asp Ser Glu His 210 215 220

Ile Ser Ser Ile Val Asp Val Pro Pro Arg Glu Ser His Ser His Ser 225 230 235 240

Asp Gln Ser Ser Lys Thr Ser Leu Met Ser Glu Met Arg Asn Ala Gln 245 250 255

Ser Ile Gly Arg Arg Trp Glu Lys Pro Ser Pro Ser Asn Val Thr Glu 260 265 270

Arg Lys Lys Arg Gly Ser Ser Trp Glu Ser Asn Asn Leu Ser Ala Asp 275 280 285

Thr Pro Cys Ala Thr Val Leu Asp Lys Gln His Ile Ser Ser Pro Glu 290 295 300

Leu Asn Cys Asn Asn Glu Ile Asn Gly His Thr Asn Glu Thr Asn Thr 305 310 315 320

- Glu Met Gln Arg Asn Lys Gln Asp Leu Pro Gly Leu Ser Ser Glu Ser 325 330 335
- Ala Arg Glu Pro Asn Ala Glu Cys Met Asn Gln Ile Glu Asp Asn Asp 340 345 350
- Asp Phe Gln Leu Gln Lys Thr Val Tyr Asp Ala Asp Met Asp Leu Thr 355 360 . 365
- Ala Ser Glu Val Ser Lys Ile Val Thr Val Ser Thr Gly Ile Lys Lys  $370 \hspace{1.5cm} 375 \hspace{1.5cm} 380$
- Lys Ser Asn Lys Lys Thr Asn Glu His Gly Met Lys Thr Phe Arg Lys 385 390 395 400
- Val Lys Asp Ser Ser Ser Glu Lys Lys Arg Glu Arg Ser Lys Arg Gln  $\cdot$  405 410 415
- Phe Lys Asn Ser Ser Asp Val Asp Ile Gly Glu Lys Ile Glu Asn Arg 420 425 430
- Thr Glu Arg Ser Asp Val Leu Asp Gly Lys Arg Gly Ala Glu Asp Pro 435 440 445
- Gly Leu Phe Phe Asn Asn Glu Gln Leu Ala Gln Met Asn Glu Gln Leu 450 460
- Ala Gln Val Asn Glu Leu Lys Lys Met Thr Leu Gln Thr Gly Phe Glu 465 470 475 480
- Gln Gly Asp Arg Glu Asn Val Leu Cys Asn Lys Lys Glu Lys Arg Val 485 490 495
- Thr Asn Glu Glu Glu Thr Tyr Ser Leu Ser Gln Ser Ser Gly Lys 500 505 510
- Phe His Gln Glu Ser Lys Phe Asp Lys Gly Gln Asn Ser Leu Thr Cys 515 520 525
- Asn Lys Ser Lys Ala Ser Arg Gln Thr Phe Val Ile His Lys Leu Glu 530  $\cdot$  540

Lys Asp Asn Leu Leu Pro Asn Gln Lys Asp Lys Val Thr Ile Tyr Glu 545 550 555 560

Asn Leu Asp Val Thr Asn Glu Phe His Thr Ala Asn Leu Ser Thr Lys 565 570 575

Asp Asn Gly Asn Leu Cys Asp Tyr Gly Thr His Asn Ile Leu Asp Leu 580 585 590

Lys Lys Tyr Val Thr Asp Ile Gln Pro Ser Glu Gln Asn Glu Ser Asn 595 600 605

Ile Asn Lys Leu Arg Lys Lys Val Asn Arg Lys Thr Glu Ile Ile Ser
610 620

Gly Met Asn His Met Tyr Glu Asp Asn Asp Lys Asp Val Val His Gly 625 630 635 640

Leu Lys Lys Gly Asn Phe Phe Phe Lys Thr Gln Glu Asp Lys Glu Pro 645  $\phantom{000}650$   $\phantom{000}655$ 

Ile Ser Glu Ser Ile Glu Val Ser Lys Glu Leu Gln Ile Pro Ala Leu 660 665 670

Ser Thr Arg Asp Asn Glu Asn Gln Cys Asp Tyr Arg Thr Gln Asn Val  $\phantom{0}675\phantom{0}$   $\phantom{0}680\phantom{0}$ 

Leu Gly Leu Gln Lys Gln Ile Thr Asn Met Tyr Pro Val Gln Gln Asn 690 695 700

Glu Ser Lys Val Asn Lys Lys Leu Arg Gln Lys Val Asn Arg Lys Thr 705 710 715 720

Glu Ile Ile Ser Glu Val Asn His Leu Asp Asn Asp Lys Ser Ile Glu 725 730 735

Tyr Thr Val Lys Ser His Ser Leu Phe Leu Thr Gln Lys Asp Lys Glu 740 745 750

Ile Ile Pro Gly Asn Leu Glu Asp Pro Ser Glu Phe Glu Thr Pro Ala 755 760 765

Leu Ser Thr Lys Asp Ser Gly Asn Leu Tyr Asp Ser Glu Ile Gln Asn

770 775 780

Val Leu Gly Val Lys His Gly His Asp Met Gln Pro Ala Cys Gln Asn 785 790 795 800

Asp Ser Lys Ile Gly Lys Lys Pro Arg Leu Asn Val Cys Gln Lys Ser 805 810 815

Glu Ile Ile Pro Glu Thr Asn Gln Ile Tyr Glu Asn Asp Asn Lys Gly 820 825 830

Val His Asp Leu Glu Lys Asp Asn Phe Phe Ser Leu Thr Pro Lys Asp 835 840 845

Lys Glu Thr Ile Ser Glu Asn Leu Gln Val Thr Asn Glu Phe Gln Thr 850 855 860

Val Asp Leu Leu Ile Lys Asp Asn Gly Asn Leu Cys Asp Tyr Asp Thr 865 870 875 880

Gln Asn Ile Leu Glu Leu Lys Lys Tyr Val Thr Asp Arg Lys Ser Ala 885 890 895

Glu Gln Asn Glu Ser Lys Ile Asn Lys Leu Arg Asn Lys Val Asn Trp 900 905 910

Lys Thr Glu Ile Ile Ser Glu Met Asn Gln Ile Tyr Glu Asp Asn Asp 915 920 925

Lys Asp Ala His Val Gln Glu Ser Tyr Thr Lys Asp Leu Asp Phe Lys 930 935 940

Val Asn Lys Ser Lys Gln Lys Leu Glu Cys Gln Asp Ile Ile Asn Lys 945 950 955 960

His Tyr Met Glu Val Asn Ser Asn Glu Lys Glu Ser Cys Asp Gln Ile 965 970 975

Leu Asp Ser Tyr Lys Val Val Lys Lys Arg Lys Lys Glu Ser Ser Cys 980 985 990

Lys Ala Lys Asn Ile Leu Thr Lys Ala Lys Asn Lys Leu Ala Ser Gln 995 1000 1005

- Leu Thr Glu Ser Ser Gln Thr Ser Ile Ser Leu Glu Ser Asp Leu 1010 1015 1020
- Lys His Ile Thr Ser Glu Ala Asp Ser Asp Pro Gly Asn Pro Val 1025 1030 1035
- Glu Leu Cys Lys Thr Gln Lys Gln Ser Thr Thr Thr Leu Asn Lys 1040 1045 1050
- Lys Asp Leu Pro Phe Val Glu Glu Ile Lys Glu Gly Glu Cys Gln 1055 1060 1065
- Val Lys Lys Val Asn Lys Met Thr Ser Lys Ser Lys Lys Arg Lys 1070 1080
- Thr Ser Ile Asp Pro Ser Pro Glu Ser His Glu Val Met Glu Arg 1085 1090 1095
- Ile Leu Asp Ser Val Gl<br/>n Gly Lys Ser Thr Val Ser Glu Gl<br/>n Ala 1100 1105 1110
- Asp Lys Glu Asn Asn Leu Glu Asn Glu Lys Met Val Lys Asn Lys 1115 1120 1125
- Pro Asp Phe Tyr Thr Lys Ala Phe Arg Ser Leu Ser Glu Ile His 1130 1135 1140
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- Asp Glu His Glu Lys Met Asn Lys Met Lys Phe Lys Val Asn Arg 1190 1195 1200
- Arg Thr Gln Lys Ser Gly Ile Gly Asp Arg Pro Leu Gln Asp Leu 1205 1210 1215

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